(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization

International Bureau



(43) International Publication Date 29 April 2004 (29.04.2004)

PCT

(10) International Publication Number WO 2004/035604 A2

(51) International Patent Classification7:

C07K

(21) International Application Number:

PCT/US2003/032812

- (22) International Filing Date: 16 October 2003 (16.10.2003)
- (25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data: 60/419,382

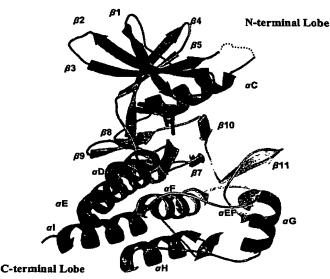
16 October 2002 (16.10.2002)

- (71) Applicant (for all designated States except US): MIL-LENNIUM PHARMACEUTICALS, INC. [US/US]; 75 Sidney Street, Cambridge, MA 02139 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): WILLIAMS, David, H. [GB/GB]; Granta Park, Great Abington, Cambridge CB1 6ET (GB). LAMERS, Maria, Bardina, Antonia, Cornelia [NL/GB]; Granta Park, Great Abington, Cambridge CB1 6ET (GB). SMITH, Stephanie [GB/GB];

- 31 Kingswood Avenue, Belper, Derbyshire DE56 1TU (GB), CHERRY, Michael [GB/GB]; Granta Park, Great Abington, Cambridge CB1 6ET (GB).
- (74) Agents: HALEY, James, F., Jr. et al.; c/o Fish & Neave, 1251 Avenue of the Americas, New York, NY 10020 (US).
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

[Continued on next page]

(54) Title: SPLEEN TYROSINE KINASE CATALYTIC DOMAIN:CRYSTAL STRUCTURE AND BINDING POCKETS THEREOF



(57) Abstract: The present invention provides crystalline molecules and molecular complexes that comprise binding pockets of Sykcat and its homologues. The invention also provides crystals comprising the catalytic domain of Syk protein. The invention further provides a computer comprising a data storage medium encoded with the structure coordinates of Sykcat binding pockets and methods for using a computer to evaluate the ability of a chemical entity or compounds to bind to a crystalline molecule or molecular complex of the invention. This invention also provides methods of using the structure coordinates to solve the structure of homologous proteins or protein complexes. The invention further provides methods of using the structure coordinates to screen for, design and optimize chemical entities or compounds, including inhibitory compounds, that bind to the catalytic domain of Syk or homologues thereof.

